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## **Search History**

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<u>L4</u>	immunosuppress\$	10829	<u>L4</u>
<u>L3</u>	11 and 12	25	<u>L3</u>
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T.1	gn19k	37	L1

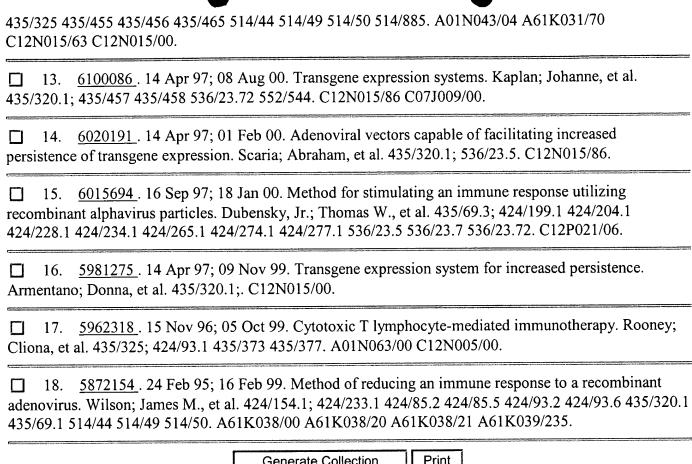
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# Search Results - Record(s) 1 through 18 of 18 returned.

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Terms	Documents
13 and 15	18

Previous Page Next Page

#### => d his (FILE 'HOME' ENTERED AT 17:08:01 ON 24 JUL 2002) FILE 'MEDLINE, CAPLUS, BIOSIS, SCISEARCH' ENTERED AT 17:08:21 ON 24 JUL 237313 S IMMUNOSUPPRESS? Ll 83 S GP19K L2L310 S L1 AND L2 4 DUP REM L3 (6 DUPLICATES REMOVED) L4330 S MHC-1 1.5 0 S L2 AND L5 L6 117157 S MHC L7 0 S L2 AND L6 L834 S L2 AND L7 L9 14 DUP REM L9 (20 DUPLICATES REMOVED) L10 => d au ti so 1-14 l10 ANSWER 1 OF 14 CAPLUS COPYRIGHT 2002 ACS Kaplan, Johanne; Armentano, Donna; Gregory, Richard J. TIAdenoviral vectors comprising a modified e4 region but retaining e4orf3 SO PCT Int. Appl., 52 pp. CODEN: PIXXD2 ANSWER 2 OF 14 CAPLUS COPYRIGHT 2002 ACS DUPLICATE 1 L10 Wold, William S. M.; Tollefson, Ann E. ΑU Adenovirus E3 proteins: 14.7K, RID, and gp19K inhibit ΤI immune-induced cell death; adenovirus death protein promotes cell death Seminars in Virology (1998), 8(6), 515-523 SO CODEN: SEVIEL; ISSN: 1044-5773 ANSWER 3 OF 14 CAPLUS COPYRIGHT 2002 ACS L10ΑU Sparer, Tim E.; Gooding, Linda R. Suppression of MHC class I antigen presentation by human ΤI adenoviruses Current Topics in Microbiology and Immunology (1998), 232 (Antigen Presentation), 135-147 CODEN: CTMIA3; ISSN: 0070-217X T<sub>1</sub>10 ANSWER 4 OF 14 MEDLINE ΑU Bruder J T; Jie T; McVey D L; Kovesdi I Expression of $\mathtt{gp19K}$ increases the persistence of transgene ΤI expression from an adenovirus vector in the mouse lung and liver. SO JOURNAL OF VIROLOGY, (1997 Oct) 71 (10) 7623-8. Journal code: 0113724. ISSN: 0022-538X. DUPLICATE 2 L10 ANSWER 5 OF 14 MEDLINE Schowalter D B; Tubb J C; Liu M; Wilson C B; Kay M A ΑU Heterologous expression of adenovirus E3-gp19K in an Ela-deleted TIadenovirus vector inhibits MHC I expression in vitro, but does not prolong transgene expression in vivo. SO GENE THERAPY, (1997 Apr) 4 (4) 351-60. Journal code: 9421525. ISSN: 0969-7128. ANSWER 6 OF 14 CAPLUS COPYRIGHT 2002 ACS Bach, Jean-Francois; Chatenoud, Lucienne; Haddada, Hedi; Lee, Martin; Perricaudet, Michel; Webb, Michelle Therapeutic gene- and immunoprotective gene-containing recombinant

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adenovirus and immunosuppressive agent medicinal combination useful for in vivo exogenic transfection and expression SO PCT Int. Appl., 40 pp. CODEN: PIXXD2 L10 ANSWER 7 OF 14 CAPLUS COPYRIGHT 2002 ACS Lee, Martin; Perricaudet, Michel IN Defective adenoviruses for gene therapy including a therapeutic gene and TIgene that protects transgenic cells from immune responses SO PCT Int. Appl., 40 pp. CODEN: PIXXD2 DUPLICATE 3 L10 ANSWER 8 OF 14 MEDLINE Basler C F; Droguett G; Horwitz M S ΑU Sequence of the immunoregulatory early region 3 and flanking sequences of TI adenovirus type 35. SO GENE, (1996 May 8) 170 (2) 249-54. Journal code: 7706761. ISSN: 0378-1119. L10 ANSWER 9 OF 14 MEDLINE DUPLICATE 4 Fejer G; Gyory I; Tufariello J; Horwitz M S ΑU Characterization of transgenic mice containing adenovirus early region 3 TIgenomic DNA. SO JOURNAL OF VIROLOGY, (1994 Sep) 68 (9) 5871-81. Journal code: 0113724. ISSN: 0022-538X. DUPLICATE 5 L10 ANSWER 10 OF 14 MEDLINE Wilson-Rawls J; Deutscher S L; Wold W S ΑU The signal-anchor domain of adenovirus E3-6.7K, a type III integral ΤI membrane protein, can direct adenovirus E3-gp19K, a type I integral membrane protein, into the membrane of the endoplasmic reticulum. VIROLOGY, (1994 May 15) 201 (1) 66-76. Journal code: 0110674. ISSN: 0042-6822. L10 ANSWER 11 OF 14 MEDLINE DUPLICATE 6 Hermiston T W; Tripp R A; Sparer T; Gooding L R; Wold W S ΑU Deletion mutation analysis of the adenovirus type 2 E3-gp19K ΤI protein: identification of sequences within the endoplasmic reticulum lumenal domain that are required for class I antigen binding and protection from adenovirus-specific cytotoxic T lymphocytes. JOURNAL OF VIROLOGY, (1993 Sep) 67 (9) 5289-98. SO Journal code: 0113724. ISSN: 0022-538X. MEDLINE DUPLICATE 7 L10 ANSWER 12 OF 14 Hermiston T W; Hellwig R; Hierholzer J C; Wold W S Sequence and functional analysis of the human adenovirus type 7 E3-ΤI gp19K protein from 17 clinical isolates. VIROLOGY, (1993 Dec) 197 (2) 593-600. SO Journal code: 0110674. ISSN: 0042-6822. **DUPLICATE 8** L10ANSWER 13 OF 14 MEDLINE Rawle F C; Tollefson A E; Wold W S; Gooding L R AU Mouse anti-adenovirus cytotoxic T lymphocytes. Inhibition of lysis by E3 ΤI gp19K but not E3 14.7K. JOURNAL OF IMMUNOLOGY, (1989 Sep 15) 143 (6) 2031-7. SO Journal code: 2985117R. ISSN: 0022-1767.

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     ANSWER 7 OF 14 CAPLUS COPYRIGHT 2002 ACS
     1996:388336 CAPLUS
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     Defective adenoviruses for gene therapy including a therapeutic gene and
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     Rhone-Poulenc Rorer S.A., Fr.
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	FΙ	9701613	Α	19970416	FI 1997-1613	19970416
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	WO	1995-FR1326	W	19951011		

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### L10 ANSWER 4 OF 14 MEDLINE

Activation of the cellular immune system and subsequent lysis of vector-transduced cells by adenovirus- or transgene-specific cytotoxic T lymphocytes have been shown to limit transgene expression in animal models. The adenovirus gp19K gene product associates with major histocompatibility complex class I proteins and prevents their maturation by sequestering them in the endoplasmic reticulum. gp19K has been shown to block the ability of adenovirus-specific cytotoxic T lymphocytes to recognize virus-infected cells in vitro. To determine if gp19K expression in an adenovirus vector would increase transgene persistence, a vector that replaces the El region of adenovirus with an expression cassette encoding both gp19K and beta-glucuronidase was constructed. This vector produced high levels of functional gp19K in infected cells. RNase protection analysis revealed efficient expression of the gp19K gene in the mouse lung. Enhanced persistence and increased beta-glucuronidase activity were observed in the lung and liver following delivery of the gp19K -expressing adenovirus vector in B10.HTG mice but not in BALB/c mice. Since gp19K binds to both class I alleles on B10.HTG mice but only one allele on BALB/c mice, these results suggest that the major histocompatibility complex class I haplotype of mice is important in determining the effectiveness of gp19K in vivo. Since gp19K has previously been shown to interact with every human major histocompatibility complex class I allele tested, the inclusion of gp19K in gene therapy vectors may increase vector persistence in human gene therapy trials.

L10 ANSWER 5 OF 14 MEDLINE An Ela-deleted adenovirus vector constitutively expressing native AB adenovirus E3-gp19K (Ad.RSV-gp19K) was constructed in order to determine whether or not E3-gp19K mediated interference with antigen presentation would result in prolonged transgene expression in vivo. Cultured fibroblasts infected with Ad.RSV-gp19K produced a native size gp19K protein and had decreased cell surface levels of MHC I as shown by immunoprecipitation and flow cytometry. The congenic mouse strains Balb/b (H-2b MHC I with high gp19K affinity), Balb/k (H-2k MHC I with no gp19K affinity), and Balb/c (H-2d MHC I with moderate gp19K affinity) were chosen for in vivo experiments because of their range of gp19K affinities. Following transduction of mice form each strain with Ad.RSV-gp19K and AD/RSV-hAAT (a reporter adenovirus), or Ad/RSV-cFIX (control adenovirus) and Ad/RSV-hAAT, the level and duration of serum hAAT protein were unrelated to gp19K protein expression. Evaluation of MHC I abundance on hepatocytes following in vivo transduction demonstrated that recombinant adenovirus rapidly increased the abundance of surface MHC I molecules on hepatocytes, and surface MHC I molecules were reduced earlier

and to a greater extent following wild-type adenovirus infection compared with hepatocytes transduced with control or Ad.RSV-gp19K recombinant adenovirus. This difference in surface MHC I down-regulation may be related to the different promoters (RSV-LTR versus the native E3 promoter) and will be an important consideration in the development of newer generation adenovirus vectors designed to evade host immune responses.

## => d bib 1 l10

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L10 ANSWER 1 OF 14 CAPLUS COPYRIGHT 2002 ACS
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    Adenoviral vectors comprising a modified e4 region but retaining e4orf3
IN
     Kaplan, Johanne; Armentano, Donna; Gregory, Richard J.
PA
     Genzyme Corp., USA
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     PCT Int. Appl., 52 pp.
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kinase. Finally, the invention provides cells transformed with said

recombinant adenoviral vectors and use of said recombinant adenoviral vectors in treatment of cancer.

L4 ANSWER 2 OF 4 MEDLINE

DUPLICATE 1

AN 2000305574 MEDLINE

DN 20305574 PubMed ID: 10845857

TI Second-generation adenoviral vectors do not prevent rapid loss of transgene expression and vector DNA from the arterial wall.

CM Comment in: Arterioscler Thromb Vasc Biol. 2000 Jun; 20(6):1414-6

AU Wen S; Schneider D B; Driscoll R M; Vassalli G; Sassani A B; Dichek D A

CS Gladstone Institute of Cardiovascular Disease, University of California, San Francisco 94141-9100, USA.

NC HL 60504 (NHLBI) P30 MH59047 (NIMH)

SO ARTERIOSCLEROSIS, THROMBOSIS, AND VASCULAR BIOLOGY, (2000 Jun) 20 (6) 1452-8.

Journal code: 9505803. ISSN: 1079-5642.

CY United States

DT Journal; Article; (JOURNAL ARTICLE)

LA English

FS Priority Journals

EM 200006

ED Entered STN: 20000706

Last Updated on STN: 20000706

Entered Medline: 20000622

AB The utility of adenoviral vectors for arterial gene transfer is limited

by

the brevity of their expression and by inflammatory host responses. As a step toward circumventing these difficulties, we used a rabbit model of

in

vivo arterial gene transfer to test 3 second-generation vectors: a vector containing a temperature-sensitive mutation in the E2A region, a vector deleted of E2A, and a vector that expresses the immunomodulatory 19-kDa glycoprotein (gp19k) from adenovirus 2. Compared with similar first-generation vectors, the second-generation vectors did not significantly prolong beta-galactosidase transgene expression or decrease inflammation in the artery wall. Although cyclophosphamide ablated the immune and inflammatory responses to adenovirus infusion, it only marginally prolonged transgene expression (94% drop in expression between 3 and 14 days). In experiments performed with "null" adenoviral vectors (no transgene), loss of vector DNA from the arterial wall was also rapid (>99% decrease between 1 hour and 14 days), unrelated to dose, and only marginally blunted by cyclophosphamide. Thus, the early loss of transgene expression after adenoviral arterial gene transfer is due primarily to loss of vector DNA, is not correlated with the presence of local vascular inflammation, and cannot be prevented by use of E2A-defective viruses, expression of gp19k, or cyclophosphamide-mediated

immunosuppression. Adenovirus-induced vascular inflammation can be
 prevented by cyclophosphamide treatment or by lowering the dose of
infused

virus. However, stabilization of adenovirus-mediated transgene expression in the arterial wall is a more elusive goal and will require novel approaches that prevent the early loss of vector DNA.

- L4 ANSWER 3 OF 4 CAPLUS COPYRIGHT 2002 ACS
- AN 1996:616318 CAPLUS
- DN 125:238676
- TI Therapeutic gene- and immunoprotective gene-containing recombinant adenovirus and immunosuppressive agent medicinal combination useful for in vivo exogenic transfection and expression

Bach, Jean-Francois; Chatenoud, Lucienne; Haddada, Hedi; Lee, Martin; IN Perricaudet, Michel; Webb, Michelle PΑ Rhone-Poulenc Rorer S.A., Fr.; Institut National De La Sante Et De La Recherche Medicale SO PCT Int. Appl., 40 pp. CODEN: PIXXD2 DT Patent French LA FAN.CNT 1 PATENT NO. KIND DATE APPLICATION NO. DATE ----------PΙ WO 9625177 A1 19960822 WO 1996-FR218 19960212 W: AL, AM, AU, BB, BG, BR, CA, CN, CZ, EE, FI, GE, HU, IS, JP, KP, KR, LK, LR, LT, LV, MD, MG, MK, MN, MX, NO, NZ, PL, RO, SG, SI, SK, TR, TT, UA, US, UZ, VN, AZ, BY, KG, KZ, RU, TJ, TM RW: KE, LS, MW, SD, SZ, UG, AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG FR 2730411 A1 19960814 FR 1995-1662 19950214 FR 2730411 B1 19970328 CA 2211039 AA 19960822 CA 1996-2211039 19960212 AU 9647238 A1 19960904 AU 1996-47238 AU 717218 B2 20000323 BR 9607310 Α 19971125 BR 1996-7310 19960212 EP 809516 Α1 19971203 EP 1996-903080 19960212 EP 809516 В1 20010822 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, PT, IE, SI JP 11500430 T219960212 19990112 JP 1996-524707 AT 204481 Ε 20010915 AT 1996-903080 19960212 ES 2163612 T3 20020201 ES 1996-903080 19960212 ZA 9601161 Α 19960807 ZA 1996-1161 19960213 FI 9703323 Α 19970813 FI 1997-3323 19970813 NO 9703724 Α 19970813 NO 1997-3724 19970813 PRAI FR 1995-1662 Α 19950214 WO 1996-FR218 W 19960212 A medicinal combination is disclosed which contains .gtoreq.1 AΒ immunosuppressive agent and .gtoreq.1 recombinant adenovirus with a genome that includes a 1st recombinant DNA contg. a therapeutic gene and 2nd recombinant DNA contg. an immunoprotective gene, for consecutive, intermittent and/or simultaneous use in in vivo and/or ex vivo exogenic transfections. The methodol. of the invention provides protection of vectors and infected cells from the immune system, thereby preventing the rapid elimination of adenovirus from infected cells and prolonging expression of the virus-carried therapeutic gene. L4ANSWER 4 OF 4 MEDLINE DUPLICATE 2 AN96235144 MEDLINE DN PubMed ID: 8666254 TISequence of the immunoregulatory early region 3 and flanking sequences of adenovirus type 35. ΑU Basler C F; Droguett G; Horwitz M S CS Department of Microbiology and Immunology, Albert Einstein College of Medicine, Bronx, NY 10461, USA. NC 5T32 CA 09060 (NCI) AI27199 (NIAID) CA-13330 (NCI) SO GENE, (1996 May 8) 170 (2) 249-54.

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CY Netherlands

DT Journal; Article; (JOURNAL ARTICLE)

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AB Adenovirus type 35 (Ad35) is an important pathogen in immunosuppressed individuals such as AIDS patients and bone marrow transplant recipients. Ad35, a member of Ad subgroup B, differs with respect to pathogenic properties from the more fully characterized subgroup C Ad, such as Ad2 and Ad5. One region of human Ad which varies between subgroups and which may influence Ad pathogenesis is early region 3 (E3), a region which appears to modulate the immune response to Ad infection. In order to begin to characterize the differences between the Ad35 E3 and the E3 of other Ad, the complete DNA sequence of the Ad35 E3 promoter and coding sequence along with two flanking structural proteins, pVIII and fiber, has been determined. Ad35 contains open reading frames which are unique to the subgroup B Ad in addition to the four characterized immunoregulatory proteins encoded by the subgroup C Ad. Further evaluation of the sequence of one of these proteins, 18.5K, which is the class-I major histocompatibility complex (MHC) binding protein of 18.5 kDa, demonstrates that the amino acid sequence of this Ad2 gp19K homologue fits a proposed model of gp19K-MHC interaction. Analysis of promoter sequences demonstrates that an NF-kappa B site found in the subgroup C E3 promoter is absent from the Ad35 E3 promoter. In addition, the fiber genes of Ad35 and other subgroup B Ad have been shown to diverge in an unexpected way, yielding three clusters of fiber homology.

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